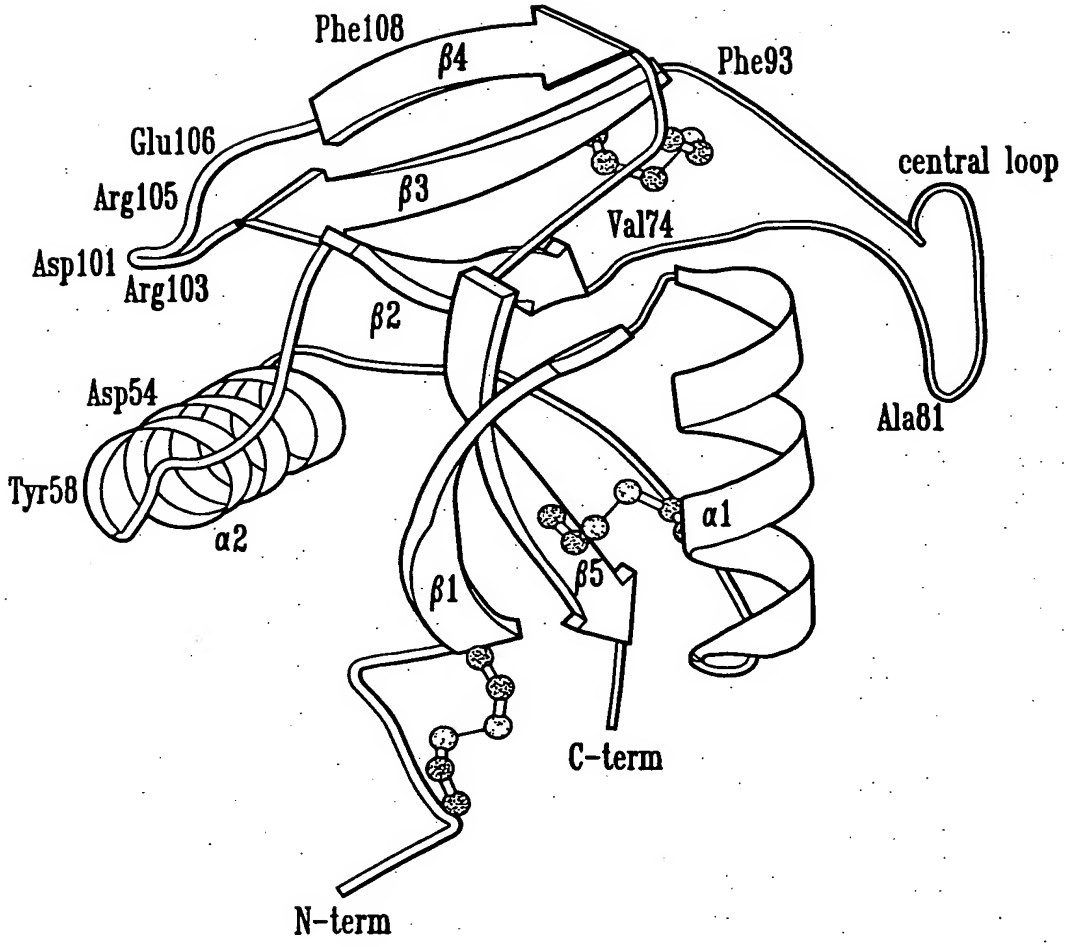
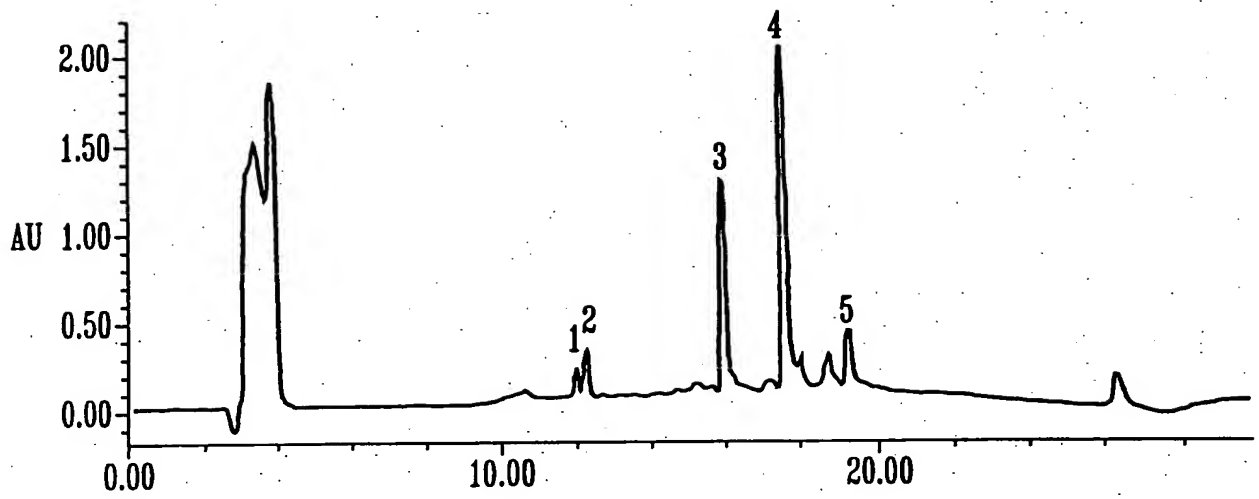
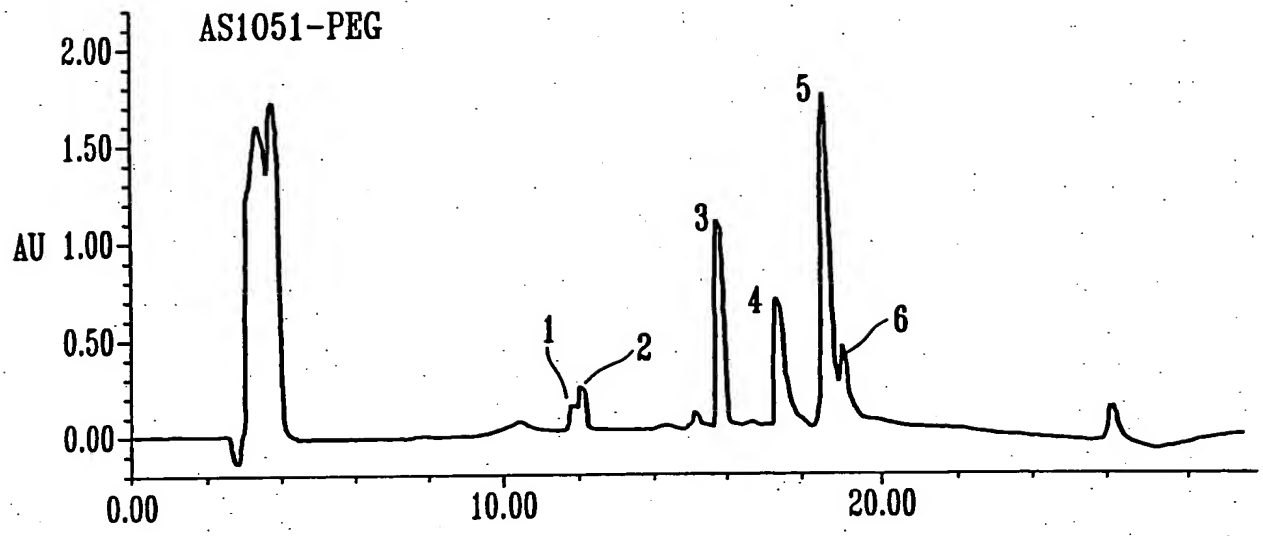

Phe108
β4
Phe93
Glu106
central loop
Arg105
β3
Val74
Asp101
Arg103
β2
Asp54
Ala81
Tyr58
α2
α1
β1
β5
C-term
N-term

FIG. 1



1 PFK
2 FTRPR
3 DLECPSGWSSYDRYCYK
QEMTWADAERFCSEQAK
4 VDCEQQHSFICK
EYLTRYIWIGLRECFMVSRDTRLREWFK
5 GGHLLSVETALEASFVDNVLYANK

FIG. 2



1 PFK

2 FTRPR

3 DLECPSGWSSYDRYCYK
QEMTWADAERFCSEQAK

4 VDCEQQHSFICK
GQPXSSISYENLVDPFECFMVSRDTRLREWFK

X=Cys-PEG

5 GGHLLSVETALEASFVDNVLYANK

FIG. 3

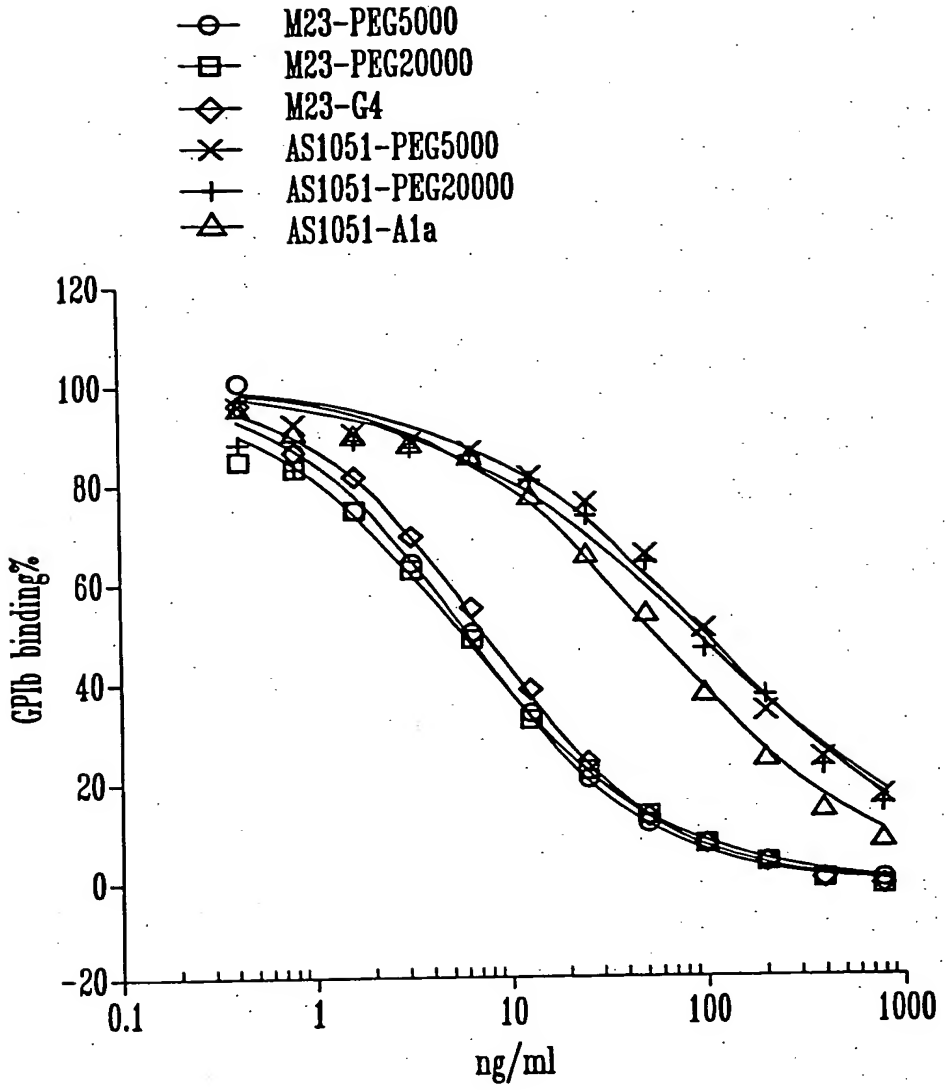

M23-PEG5000
M23-PEG20000
M23-G4
AS1051-PEG5000
AS1051-PEG20000
AS1051-A1a
GPIb binding%
120
100
80
60
40
20
0
-20
0.1
1
10
100
1000
ng/ml

FIG. 4

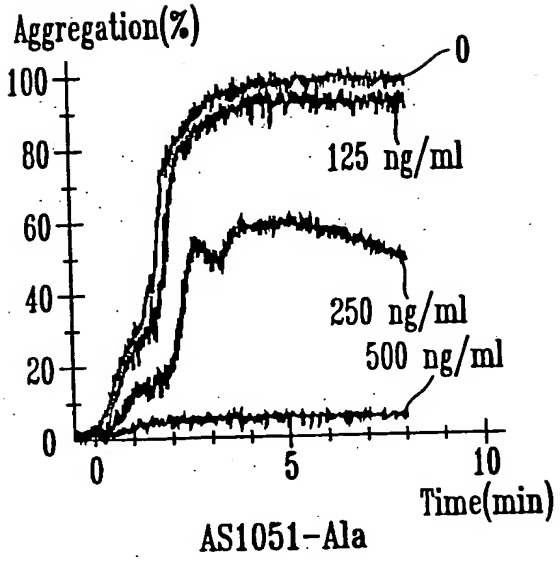


FIG. 5A

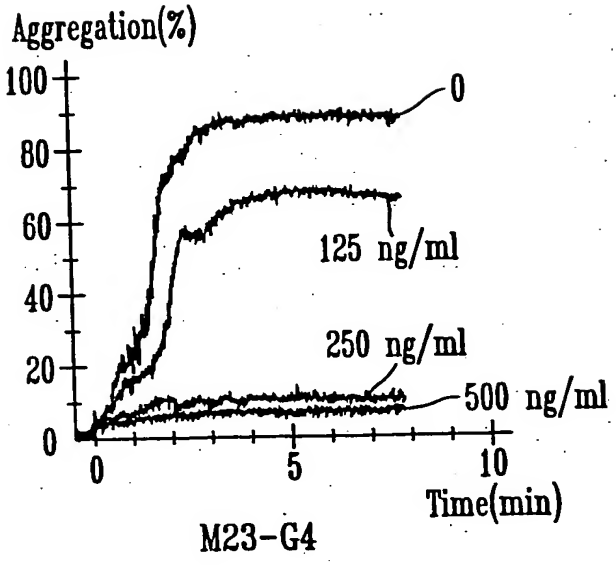


FIG. 5B

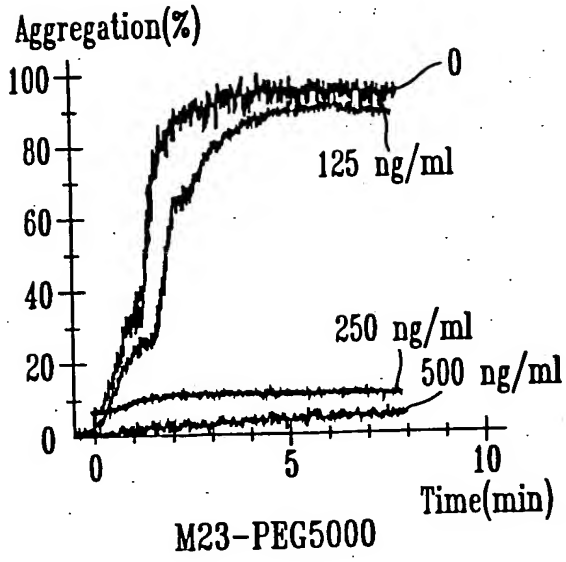


FIG. 5C

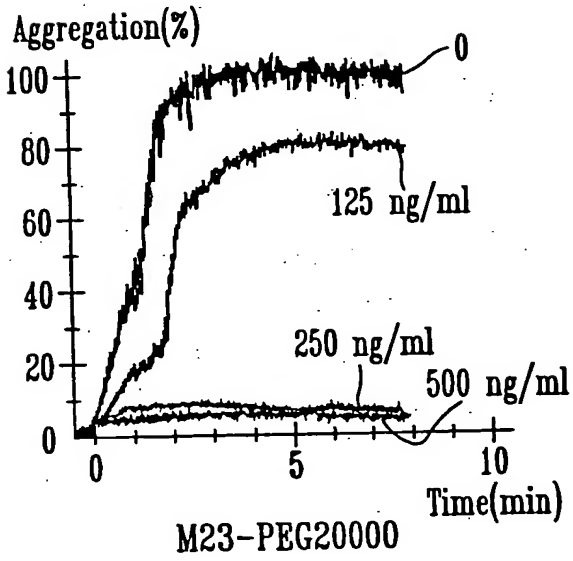


FIG. 5D

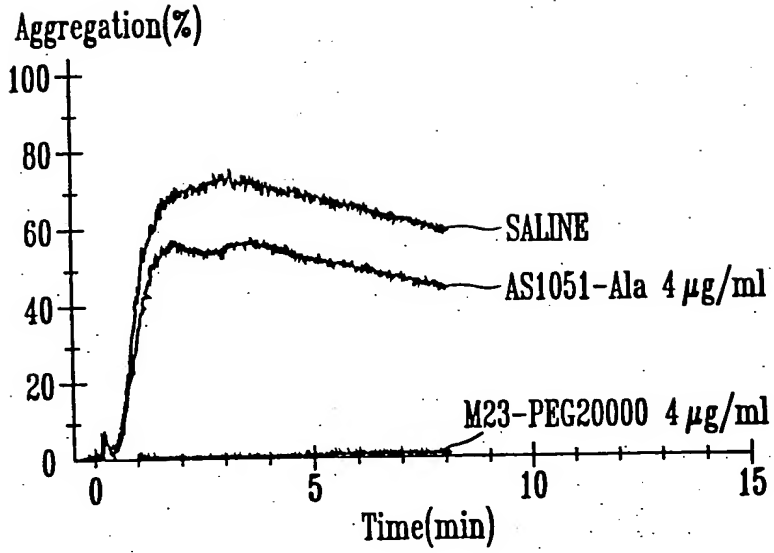

Aggregation(%)
100
80
60
40
20
0
0
5
10
15
Time(min)
SALINE
AS1051-Ala 4 μg/ml
M23-PEG20000 4 μg/ml

FIG. 6A

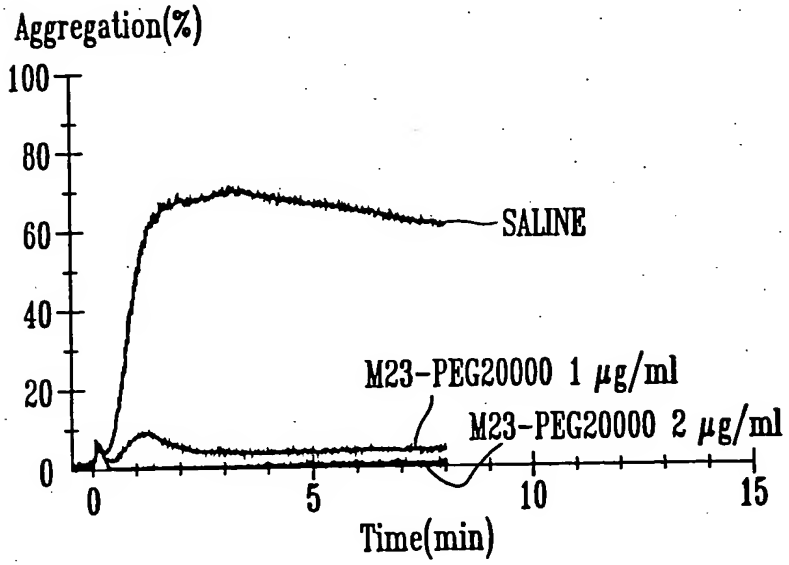

Aggregation(%)
100
80
60
40
20
0
0
5
10
15
Time(min)
SALINE
M23-PEG20000 1 μg/ml
M23-PEG20000 2 μg/ml

FIG. 6B

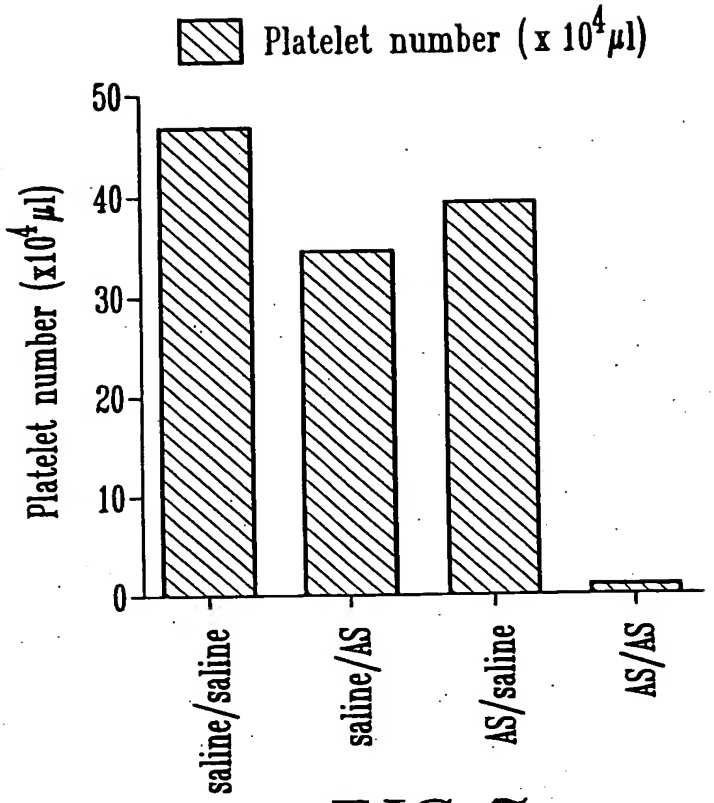

Platelet number (x 10^4 µl)
Platelet number (x10^4 µl)
50
40
30
20
10
0
saline/saline
saline/AS
AS/saline
AS/AS

FIG. 7

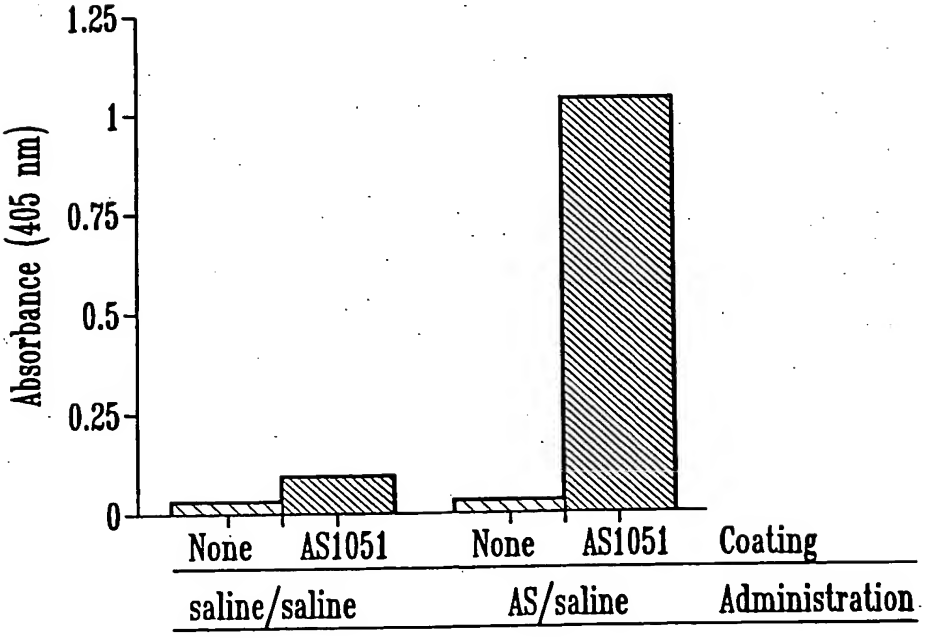

Absorbance (405 nm)
1.25
1
0.75
0.5
0.25
0
None
AS1051
None
AS1051
Coating
saline/saline
AS/saline
Administration

FIG. 8

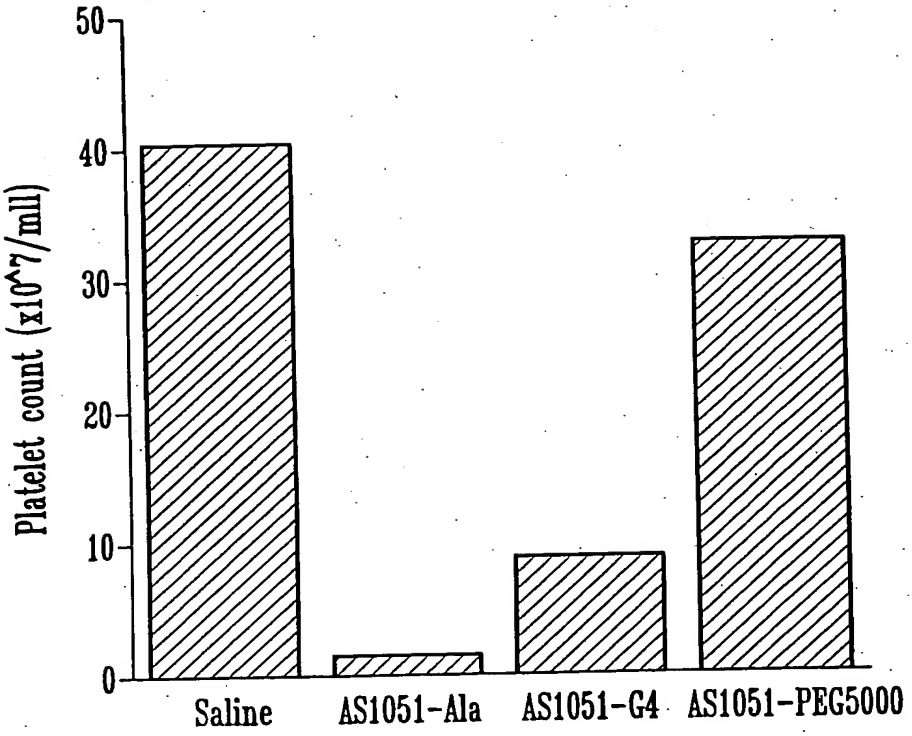

Platelet count (x10^7/ml)
50
40
30
20
10
0
Saline
AS1051-Ala
AS1051-G4
AS1051-PEG5000

FIG. 9

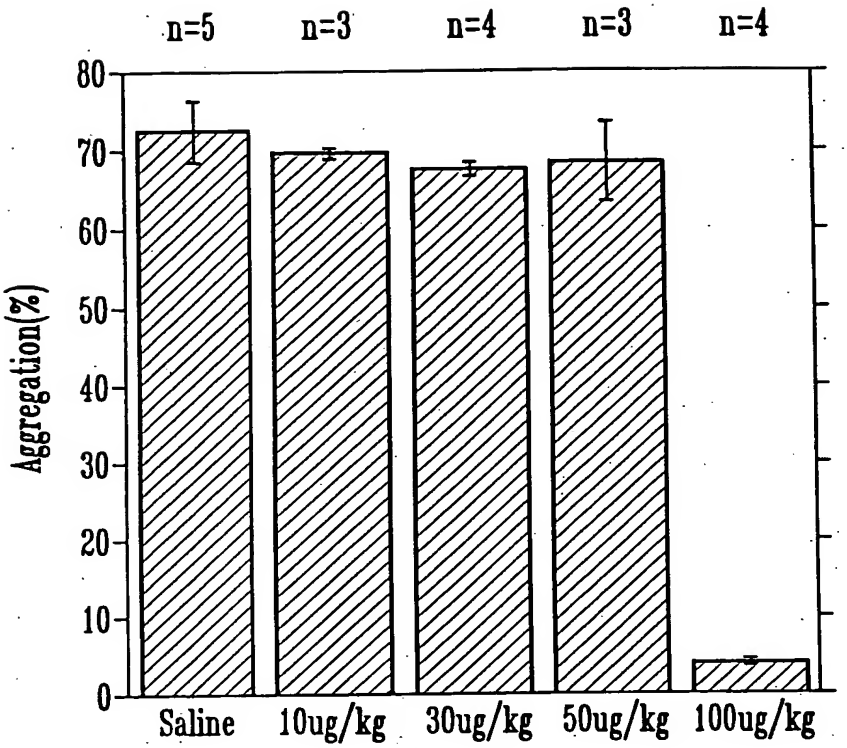


FIG. 10A

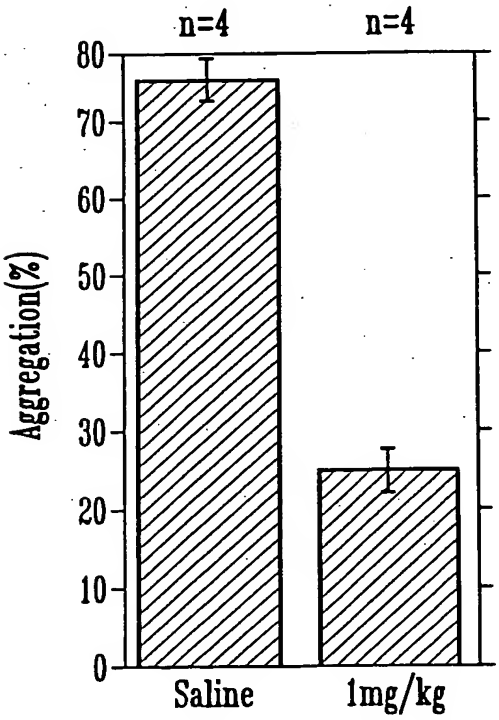


FIG. 10B